

SEQUENCE LISTING

<110> BLANC, Veronique

THIBAUT, Denis

BAMAS-JACQUES, Nathalie

BLANCHE, Francis

COUZET, Joel

BARRIERE, Jean-Claude

DEBUSSCHE, Laurent

FAMECHON, Alain

PARIS, Jean-Marc

DUTRUC-ROSSET, Gilles

<120> Streptogramins And Method For Preparing Same By Mutasyntesis

<130> Streptogramin genes

<140> 08/765,907

<141> 1997-03-20

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<170> PatentIn Ver. 2.0

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<213> *Streptomyces pristinaespiralis*

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<212> PRT

<213> Streptomyces pristinaespiralis

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Ala Val Thr Trp Leu Asp Val Ala Gly Ala Gly Ala Ala Asp Gly Val
35 40 45

Arg Val Val Ala Gly Asp Val Arg Arg Pro Gly Pro Glu Ala Val Ala
50 55 60

Ala Leu Ala Ala Ala Asp Val Val Val Leu Ala Val Pro Glu Pro Val
65 70 75 80

Ala Trp Glu Ala Val Glu Val Leu Ala Gly Val Met Arg Pro Gly Ala
85 90 95

Val Leu Ala Asp Thr Leu Ser Val Lys Ser Arg Ile Ala Gly Arg Leu
100 105 110

Arg Glu Ala Ala Pro Gly Leu Gln Ala Val Gly Leu Asn Pro Met Phe
115 120 125

Ala Pro Ser Leu Gly Leu Gln Gly Arg Pro Val Ala Ala Val Val Val
130 135 140

Thr Asp Gly Pro Gly Val Arg Ala Leu Val Glu Leu Val Ala Gly Trp
145 150 155 160

Gly Ala Arg Val Val Glu Met Pro Ala Arg Arg His Asp Glu Leu Thr
165 170 175

Ala Ala Gln Gln Ala Ala Thr His Ala Ala Val Leu Ala Phe Gly Leu
180 185 190

Gly Leu Gly Glu Leu Ser Val Asp Val Gly Ala Leu Arg Asp Ser Ala
195 200 205

Pro Pro Pro His Leu Ala Met Leu Ala Leu Leu Ala Arg Ile Ala Gly
210 215 220

Table 1. Demographic characteristics of the study population	
Age (years)	50.0 ± 10.0
Gender	
Male	50.0%
Female	50.0%
Education (years)	12.0 ± 2.0
Marital status	
Married	80.0%
Single	20.0%
Occupation	
Professional	30.0%
Managerial	20.0%
Technical	10.0%
Service	20.0%
Unemployed	20.0%
Income (USD/month)	1000.0 ± 500.0
Health status	
Good	70.0%
Fair	20.0%
Poor	10.0%

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<212> PRT

<213> Streptomyces pristinaespiralis

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 35 40 45
 Arg Ile Gly Glu Tyr Lys Arg Leu His Gln Val Pro Met Met Gln Pro
 50 55 60
 His Arg Ile Ala Gln Val His Ala Asn Ala Ala Arg Tyr Ala Ala Asp
 65 70 75 80
 His Gly Ile Asp Pro Ala Phe Leu Arg Thr Leu Tyr Asp Thr Ile Ile
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Thr Glu Thr Cys Arg Leu Glu Asp Glu Trp Ile Ala Ser Gly Gly Ala
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Ser Pro
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<211> 4496

<212> DNA

<213> *Streptomyces pristinaespiralis*

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<213> *Streptomyces pristinaespiralis*

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<212> PRT

<213> Streptomyces pristinaespiralis

<400> 8

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Met Pro His Arg Glu Pro Gly Asp His Ile Thr Leu Lys Thr Val Gly
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Tyr Ser Pro Ala Asn Pro Gly Arg Phe Gly Leu Pro Thr Ile Leu Gly
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Thr Val Ala Arg Tyr Asp Asp Thr Thr Gly Ala Leu Thr Ala Leu Met
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Asp Gly Val Leu Leu Thr Ala Leu Arg Thr Gly Ala Ala Ser Ala Val
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Ala Ser Arg Leu Leu Ala Arg Pro Asp Ser His Thr Leu Gly Leu Ile
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Gly Thr Gly Ala Gln Ala Val Thr Gln Leu His Ala Leu Ser Leu Val
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Leu Pro Leu Gln Arg Ala Leu Val Trp Asp Thr Asp Pro Ala His Arg
165 170 175

Glu Ser Phe Ala Arg Arg Ala Ala Phe Thr Gly Val Ser Val Glu Ile
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Ala Glu Pro Ala Arg Ile Ala Ala Glu Ala Asp Val Ile Ser Thr Ala
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Thr Ser Val Ala Val Gly Gln Gly Pro Val Leu Pro Asp Thr Gly Val
210 215 220

Arg Glu His Leu His Ile Asn Ala Val Gly Ala Asp Leu Val Gly Lys
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Thr Glu Leu Pro Leu Gly Leu Leu Glu Arg Ala Phe Val Thr Ala Asp
245 250 255

His Pro Glu Gln Ala Leu Arg Glu Gly Glu Cys Gln Gln Leu Ser Ala
260 265 270

Asp Arg Leu Gly Pro Gln Leu Ala His Leu Cys Ala Asp Pro Ala Ala
275 280 285

Ala Ala Gly Arg Gln Asp Thr Leu Ser Val Phe Asp Ser Thr Gly Phe
290 295 300

Ala Phe Glu Asp Ala Leu Ala Met Glu Val Phe Leu Glu Ala Ala Ala
305 310 315 320

Glu Arg Asp Leu Gly Ile Arg Val Gly Ile Glu His His Pro Gly Asp
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<213> *Streptomyces pristinaespiralis*

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<213> Streptomyces pristinaespiralis

<400> 10

Met	Pro	Pro	Thr	Pro	Arg	Pro	Thr	Thr	Asp	Asp	Gly	Gly	Arg	Glu	Leu
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Leu	Ala	Trp	Leu	Arg	Glu	Met	Arg	His	His	His	Pro	Val	His	Glu	Asp
			20					25					30		

Glu Tyr Gly Ala Phe His Val Phe Arg His Ala Asp Val Leu Thr Val
35 40 45

Ala Ser Asp Pro Gly Val Tyr Ser Ser Gln Leu Ser Arg Leu Arg Pro
50 55 60

Gly Ser Gln Ala Leu Ser Glu Gln Ile Leu Ser Val Ile Asp Pro Pro
65 70 75 80

Met His Arg Thr Leu Arg Arg Leu Val Ser Gln Ala Phe Thr Pro Arg
85 90 95

Thr Val Ala Asp Leu Glu Pro Arg Val Thr Glu Leu Ala Gly Gln Leu
100 105 110

Leu Asp Ala Val Asp Gly Asp Thr Phe Asp Leu Val Ala Asp Phe Ala
115 120 125

Tyr Pro Leu Pro Val Ile Val Ile Ala Glu Leu Leu Gly Val Pro Pro
130 135 140

Ala Asp Arg Thr Leu Phe Arg Ser Trp Ser Asp Arg Met Leu Gln Met
145 150 155 160

Gln Val Ala Asp Pro Ala Asp Met Gln Phe Gly Asp Asp Ala Asp Glu
165 170 175

Asp Tyr Gln Arg Leu Val Lys Glu Pro Met Arg Ala Met His Ala Tyr
180 185 190

Leu His Asp His Val Thr Asp Arg Arg Ala Arg Pro Ala Asn Asp Leu
195 200 205

Ile Ser Ala Leu Val Ala Ala Arg Val Glu Gly Glu Arg Leu Thr Asp
210 215 220

Glu Gln Ile Val Glu Phe Gly Ala Leu Leu Leu Met Ala Gly His Val
225 230 235 240

Ser Thr Ser Met Leu Leu Gly Asn Thr Val Leu Cys Leu Lys Asp His
245 250 255

Pro Arg Ala Glu Ala Ala Ala Arg Ala Asp Arg Ser Leu Ile Pro Ala
260 265 270

Leu Ile Glu Glu Val Leu Arg Leu Arg Pro Pro Ile Thr Val Met Ala
275 280 285

Arg Val Thr Thr Lys Asp Thr Val Leu Ala Gly Thr Thr Ile Pro Ala
290 295 300

Gly Arg Met Val Val Pro Ser Leu Leu Ser Ala Asn His Asp Glu Gln
305 310 315 320

Val Phe Thr Asp Pro Asp His Leu Asp Leu Ala Arg Glu Gly Arg Gln
325 330 335

Ile Ala Phe Gly His Gly Ile His Tyr Cys Leu Gly Ala Pro Leu Ala
340 345 350

Arg Leu Glu Gly Arg Ile Ala Leu Glu Ala Leu Phe Asp Arg Phe Pro
355 360 365

Asp Phe Ser Pro Thr Asp Gly Ala Lys Leu Arg Tyr His Arg Asp Gly
370 375 380

Leu Phe Gly Val Lys Asn Leu Pro Leu Thr Val Arg Arg Gly Pro
385 390 395

<210> 11

<211> 1561

<212> DNA

<213> *Streptomyces pristinaespiralis*

<400> 11

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tgggcgaact ggtccgctcg ggccggtcgc tgacgtcggg ggtgtggcgg gagcggttcg 360
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gcggccggca ggtgcgggag cgttcgtgc gcgccctgga ccggtgggt gtggaggtcc 1260
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 agggccacgg cacggggacg cgggtggcgg ccgggcacct gtgacaccgt ccgcatccgg 1500
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<210> 12

<211> 1233

<212> DNA

<213> *Streptomyces pristinaespiralis*

<400> 12

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 ttgccccg tgaccggcgc ccggcacgcg ctcaagtca ccagcggcac cgtcgcgtg 180
 gaactggcgg tgcggatgct ggacctggcg ccgggcgacg aggtgatcgc caccgcag 240
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 atcgaccgg acacctcaa cctcgaccg gcggtgctgg agacgtgat caccgaccgc 360
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gccagcggg tgggacgtat ccgcgacaac gaggccgacg gcgtgtacgc ggcgctgccg 660
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gcggcggcgg tgggcgtggt gcaactggcg tcgctggagc ggttcgtggc ccggcggccg 840
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<210> 13

<211> 412

<212> PRT

<213> Streptomyces pristinaespiralis

<400> 13

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Ala Leu Gly Glu Leu Val Arg Ser Gly Arg Ser Leu Thr Ser Gly Val
20 25 30

Trp Arg Glu Arg Phe Glu Glu Gln Phe Ala Arg Leu Thr Gly Ala Arg
35 40 45

His Ala Leu Ser Val Thr Ser Gly Thr Val Ala Leu Glu Leu Ala Val
50 55 60

Arg Met Leu Asp Leu Ala Pro Gly Asp Glu Val Ile Ala Thr Pro Gln
65 70 75 80

Thr Phe Gln Ala Thr Val Gln Pro Leu Leu Asp His Asp Val Arg Leu
85 90 95

Arg Phe Cys Asp Ile Asp Pro Asp Thr Leu Asn Leu Asp Pro Ala Val
100 105 110

Leu Glu Thr Leu Ile Thr Asp Arg Thr Arg Ala Ile Leu Leu Val His
115 120 125

Tyr Gly Gly Asn Pro Ala Asp Met Asp Arg Ile Met Ala Leu Ala Arg
130 135 140

Lys Arg Gly Ile Ile Val Val Glu Asp Ser Ala His Ala Leu Gly Ala
145 150 155 160

Val Tyr Arg Gly Arg Arg Pro Gly Ala Leu Ala Asp Ile Gly Cys Phe
165 170 175

Thr Phe His Ser Thr Lys Asn Ile Thr Thr Leu Gly Glu Gly Gly Met
180 185 190

Ile Thr Leu Ser Arg Asp Glu Trp Ala Gln Arg Val Gly Arg Ile Arg
195 200 205

Asp Asn Glu Ala Asp Gly Val Tyr Ala Ala Leu Pro Asp Ser Ala Arg
210 215 220

Ala Gly Ala Pro Ala Leu Leu Pro Trp Met Lys Phe Ala Glu Gly Val
225 230 235 240

Tyr Gly His Arg Ala Val Gly Val Arg Gly Ala Gly Thr Asn Ala Thr
245 250 255

Met Ser Glu Ala Ala Ala Ala Val Gly Val Val Gln Leu Ala Ser Leu
260 265 270

Glu Arg Phe Val Ala Arg Arg Arg Ser Ile Ala Gln Arg Leu Asp Glu
275 280 285

Ala Val Ala Ser Val Ala Gly Thr Arg Leu His Arg Ala Ala Ala Asp
290 295 300

Ser Leu His Ala Tyr His Leu Tyr Thr Phe Phe Leu Thr Gly Gly Arg
305 310 315 320

Gln Val Arg Glu Arg Phe Val Arg Ala Leu Asp Arg Leu Gly Val Glu
325 330 335

Val Gln Leu Arg Tyr Phe Pro Leu His Leu Ser Pro Glu Trp Arg Leu
340 345 350

Arg Gly His Gly Pro Gly Glu Cys Pro Thr Ala Glu Arg Val Trp Phe
355 360 365

Glu Glu His Met Asn Leu Pro Cys His Pro Gly Leu Ser Asp Gly Gln
370 375 380

Val Asp Tyr Met Val Glu Ala Val Thr Arg Ala Leu His Glu Ala His
385 390 395 400

Gly Thr Gly Thr Arg Val Ala Ala Gly His Leu Pro
405 410

<210> 14

<211> 2220

<212> DNA

<213> *Streptomyces pristinaespiralis*

<400> 14

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<210> 15

<211> 719

<212> PRT

<213> Streptomyces pristinaespiralis

<400> 15

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Tyr Asn Leu Phe Gln Met Leu Ala Glu Val Asn Gly Ala Ala Pro Leu
      20             25             30
Val Val Arg Asn Asp Asp Thr Arg Thr Trp Gln Ala Leu Ala Pro Gly
      35             40             45
Asp Phe Asp Asn Val Val Val Ser Pro Gly Pro Gly His Pro Ala Thr
      50             55             60
Asp Thr Asp Leu Gly Leu Ser Arg Arg Val Ile Thr Glu Trp Asp Leu
65             70             75             80
Pro Leu Leu Gly Val Cys Leu Gly His Gln Ala Leu Cys Leu Leu Ala
      85             90             95
Gly Ala Ala Val Val His Ala Pro Glu Pro Phe His Gly Arg Thr Ser
      100            105            110
Asp Ile Arg His Asp Gly Gln Gly Leu Phe Ala Asn Ile Pro Ser Pro
      115            120            125
Leu Thr Val Val Arg Tyr His Ser Leu Thr Val Arg Gln Leu Pro Ala
      130            135            140
Asp Leu Arg Ala Thr Ala His Thr Ala Asp Gly Gln Leu Met Ala Val
145            150            155            160
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Ala His Arg His Leu Pro Arg Phe Gly Val Gln Phe His Pro Glu Ser
165 170 175

Ile Ser Ser Glu His Gly His Arg Met Leu Ala Asn Phe Arg Asp Leu
180 185 190

Ser Leu Arg Ala Ala Gly His Arg Pro Pro His Thr Glu Arg Ile Pro
195 200 205

Ala Pro Ala Pro Ala Pro Ala Pro Ala Pro Ala Pro Ala Pro Pro Ala
210 215 220

Ser Ala Pro Val Gly Glu Tyr Arg Leu His Val Arg Glu Val Ala Cys
225 230 235 240

Val Pro Asp Ala Asp Ala Ala Phe Thr Ala Leu Phe Ala Asp Ala Pro
245 250 255

Ala Arg Phe Trp Leu Asp Ser Ser Arg Val Glu Pro Gly Leu Ala Arg
260 265 270

Phe Thr Phe Leu Gly Ala Pro Ala Gly Pro Leu Gly Glu Gln Ile Thr
275 280 285

Tyr Asp Val Ala Asp Arg Ala Val Arg Val Lys Asp Gly Ser Gly Gly
290 295 300

Glu Thr Arg Arg Pro Gly Thr Leu Phe Asp His Leu Glu His Glu Leu
305 310 315 320

Ala Ala Arg Ala Leu Pro Ala Thr Gly Leu Pro Phe Glu Phe Asn Leu
325 330 335

Gly Tyr Val Gly Tyr Leu Gly Tyr Glu Thr Lys Ala Asp Ser Gly Gly
340 345 350

Glu Asp Ala His Arg Gly Glu Leu Pro Asp Gly Ala Phe Met Phe Ala
355 360 365

Asp Arg Met Leu Ala Leu Asp His Glu Gln Gly Arg Ala Trp Leu Leu
370 375 380

Ala Leu Ser Ser Thr Arg Arg Pro Ala Thr Ala Pro Ala Ala Glu Arg
385 390 395 400

Trp Leu Thr Asp Ala Ala Arg Thr Leu Ala Thr Thr Ala Pro Arg Pro
 405 410 415
 Pro Phe Thr Leu Leu Pro Asp Asp Gln Leu Pro Ala Leu Asp Val His
 420 425 430
 Tyr Arg His Ser Leu Pro Arg Tyr Arg Glu Leu Val Glu Glu Cys Arg
 435 440 445
 Arg Leu Ile Thr Asp Gly Glu Thr Tyr Glu Val Cys Leu Thr Asn Met
 450 455 460
 Leu Arg Val Pro Gly Arg Ile Asp Pro Leu Thr Ala Tyr Arg Ala Leu
 465 470 475 480
 Arg Thr Val Ser Pro Ala Pro Tyr Ala Ala Tyr Leu Gln Phe Pro Gly
 485 490 495
 Ala Thr Val Leu Ser Ser Ser Pro Glu Arg Phe Leu Arg Ile Gly Ala
 500 505 510
 Asp Gly Trp Ala Glu Ser Lys Pro Ile Lys Gly Thr Arg Pro Arg Gly
 515 520 525
 Ala Gly Pro Ala Gln Asp Ala Ala Val Lys Ala Ser Leu Ala Ala Ala
 530 535 540
 Glu Lys Asp Arg Ser Glu Asn Leu Met Ile Val Asp Leu Val Arg Asn
 545 550 555 560
 Asp Leu Gly Gln Val Cys Asp Ile Gly Ser Val His Val Pro Gly Leu
 565 570 575
 Phe Glu Val Glu Thr Tyr Ala Thr Val His Gln Leu Val Ser Thr Val
 580 585 590
 Arg Gly Arg Leu Ala Ala Asp Val Ser Arg Pro Arg Ala Val Arg Ala
 595 600 605
 Ala Phe Pro Gly Gly Ser Met Thr Gly Ala Pro Lys Val Arg Thr Met
 610 615 620
 Gln Phe Ile Asp Arg Leu Glu Lys Gly Pro Arg Gly Val Tyr Ser Gly
 625 630 635 640

Ala Leu Gly Tyr Phe Ala Leu Ser Gly Ala Ala Asp Leu Ser Ile Val
645 650 655

Ile Arg Thr Ile Val Ala Thr Glu Glu Ala Ala Thr Ile Gly Val Gly
660 665 670

Gly Ala Val Val Ala Leu Ser Asp Pro Asp Asp Glu Val Arg Glu Met
675 680 685

Leu Leu Lys Ala Gln Thr Thr Leu Ala Ala Leu Arg Gln Ala His Ala
690 695 700

Gly Ala Thr Ala Ser Asp Arg Glu Leu Leu Ala Gly Ser Leu Arg
705 710 715

<210> 16

<211> 962

<212> DNA

<213> *Streptomyces pristinaespiralis*

<400> 16

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acgaggccac cgggcagctg accggcgccg ggatcaccgc cgacgccgcc cgggccgaca 180

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<210> 17

<211> 292

<212> PRT

<213> Streptomyces pristinaespiralis

<400> 17

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20	25	30	
Thr Arg Leu Leu Ala Ala His Ala Cys Gln Val Ala Pro Gly Asp Leu			
35	40	45	
Asp Thr Cys Leu Ala Gly Pro Val Pro Pro Arg Phe Trp His Tyr Val			
50	55	60	
Arg Arg Arg Leu Thr Arg Glu Pro Ala Glu Arg Ile Val Gly His Ala			
65	70	75	80
Tyr Phe Met Gly His Arg Phe Asp Leu Ala Pro Gly Val Phe Val Pro			
85	90	95	
Lys Pro Glu Thr Glu Glu Ile Thr Arg Asp Ala Ile Ala Arg Leu Glu			
100	105	110	
Ala Leu Val Arg Arg Gly Thr Thr Ala Pro Leu Val Val Asp Leu Cys			
115	120	125	
Ala Gly Pro Gly Thr Met Ala Val Thr Leu Ala Arg His Val Pro Ala			
130	135	140	
Ala Arg Val Leu Gly Ile Glu Leu Ser Gln Ala Ala Ala Arg Ala Ala			
145	150	155	160
Arg Arg Asn Ala Arg Gly Thr Gly Ala Arg Ile Val Gln Gly Asp Ala			
165	170	175	
Arg Asp Ala Phe Pro Glu Leu Ser Gly Thr Val Asp Leu Val Val Thr			
180	185	190	
Asn Pro Pro Tyr Ile Pro Ile Gly Leu Arg Thr Ser Ala Pro Glu Val			
195	200	205	
Leu Glu His Asp Pro Pro Leu Ala Leu Trp Ala Gly Glu Glu Gly Leu			
210	215	220	
Gly Met Ile Arg Ala Met Glu Arg Thr Ala Ala Arg Leu Leu Ala Pro			
225	230	235	240

Gly Gly Val Leu Leu Leu Glu His Gly Ser Tyr Gln Leu Ala Ser Val
245 250 255

Pro Ala Leu Phe Arg Ala Thr Gly Arg Trp Ser His Ala Ser Ser Arg
260 265 270

Pro Thr Cys Asn Asp Gly Cys Leu Thr Ala Val Arg Asn His Thr Cys
275 280 285

Ala Pro Pro Ala
290

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